PATENT

Attorney Docket No.: VOSS1110

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## Amendments to the Claims:

Please amend claims 1, 3, 7, and 9 as indicated in the Listing of Claims.

## **Listing of Claims:**

This listing of claims will replace all prior versions, and listings, of claims in the application:

- 1. (Currently Amended): A recombinant DNA molecule comprising:
- (a) at least one first regulatory sequence which confers expression in endothelial cells *in vivo*, wherein said first regulatory sequence is selected from the group consisting of
  - (i) a DNA sequence comprising a nucleotide sequence as given in SEQ ID NO: 1;
  - (ii) a DNA sequence comprising a nucleotide sequence of SEQ ID NO:1 from nucleotide 8260 to nucleotide 10560, from nucleotide 8336 to nucleotide10608 and/or from nucleotide 10094 to nucleotide 10608; and
  - (iii) a DNA sequence comprising a fragment of a nucleotide sequence of (i) or (ii) SEQ ID NO: 1, wherein the fragment confers endothelial cell-specific expression; and
  - (b) operatively linked thereto a heterologous DNA sequence.
- 2. (Original): The recombinant DNA molecule of claim 1, wherein said first regulatory sequence comprises a GATA-binding site, an AP-1 binding site, an SP1 binding site, an NFkB binding site, a STAT binding site, a ScI/tal-1 binding site, an Ets-1 binding site, a PEA3 consensus sequence or any combination(s) thereof
- 3. (Currently Amended) The recombinant DNA molecule of claim 1 or 2, wherein said the first regulatory sequence is a DNA sequence comprising a fragment of a nucleotide sequence from nucleotide 8260 to nucleotide 10560 of SEQ ID NO:1, from nucleotide 8336 to

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nucleotide 10608 of SEQ ID NO:1, and/or from nucleotide 10094 to nucleotide 10608 of SEQ ID NO:1, wherein the fragment confers endothelial cell-specific expression

ID NO:1, wherein the fragment confers endothelial cell-specific expression
selected from the group consisting of
(a) a DNA sequence comprising a nucleotide sequence as given in SEQ
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(b) a DNA sequence comprising a nucleotide sequence of SEQ ID NO: 1
from nucleotide 8260 to nucleotide 10560, from nucleotide 8336 to nucleotide 10608
and/or from nucleotide 10094 to nucleotide 10608; and
(c) a DNA sequence comprising a fragment of a nucleotide sequence of any
one of (a) or (b) that confers expression in endothelial cells.

- 4. (Previously Presented): The recombinant DNA molecule of any one of claims 1 to 2, wherein said heterologous DNA sequence is operatively linked to further regulatory sequences.
- 5. (Original): The recombinant DNA molecule of claim 4, wherein said further regulatory sequence is a promoter.
- 6. (Previously Presented): The recombinant DNA molecule of claim 4, wherein said further regulatory sequence is a 3'-untranslated region.
- 7. (Currently Amended): The recombinant DNA molecule of claim 5, wherein said promoter is a promoter of <u>a</u> hypoxia inducible genes gene, genes <u>a gene</u> encoding <u>a</u> growth factors factor or its receptor or <u>a</u> glycolytic enzymes enzyme.
- 8. (Original): The recombinant DNA molecule of claim 7, wherein said growth factor is VEGF, PDGF or Fibroblast growth factor.

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(Currently Amended): The recombinant DNA molecule of claim 5, wherein 9. said promoter comprises a DNA sequence selected from the group consisting of

- a DNA sequence comprising the nucleotide sequence as given in SEQ (a) ID NO: 1 from nucleotide 6036 to nucleotide 6959;
- a DNA sequence comprising the nucleotide sequence of the human Flk-1/KDR promoter; and
- a DNA sequence comprising a fragment of a nucleotide sequence of any one of (a) or (b) nucleotide 6036 to nucleotide 6959 of SEQ ID NO:1 or the nucleotide sequence of the human Flk-1/KDR promoter, wherein the fragment confers endothelial cell-specific expression.
- (Previously Presented): The recombinant DNA molecule of any one of claims 1 10. to 2, wherein at least one of said DNA sequences is of human or murine origin.
- (Previously Presented): The recombinant DNA molecule of any one of claims 1 11. to 2, wherein said heterologous DNA sequence being operatively linked to said regulatory sequences is located 5' to said first regulatory sequence.

## 12. (Canceled)

(Previously Presented): The recombinant DNA molecule of claim 42, wherein 13. said protein is selected from the group consisting of Vascular Endothelial Growth Factor (VEGF), Hypoxia Inducible Factors (HIF), HIF-Related Factor (HRF), tissue plasminogen activator, p21 cell cycle inhibitor, nitric oxide synthase, interferon-γ, atrial natriuretic polypeptide, monocyte chemotactic proteins, luciferase, green fluorescent protein and lacZ.

Claims 14 –16 (Canceled)

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17. (Previously Presented): A vector comprising a recombinant DNA molecule of any one of claims 1 to 2.

18. The vector of claim 17, which is an expression vector and/or a targeting vector.

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- 19. (Previously amended) The vector of claim 17, further comprising a gene capable of expressing HIF- $2\alpha$ .
- 20. (Previously amended) An isolated cell transformed with a DNA molecule of any one of claims 1 to 2.
- 21. (Previously amended) The isolated cell of claim 20, which is a prokaryotic or eukaryotic cell.
- 22. (Previously amended) The isolated cell of claim 20, which is an endothelial cell.
- 23. (Previously amended) The isolated cell of claim 20, further comprising a recombinant DNA molecule or vector containing a gene capable of expressing HIF- $2\alpha$ .

Claims 24-41 (Canceled)

42. (Previously Presented): The recombinant DNA molecule of any one of claims 1 to 2, wherein said heterologous DNA sequence encodes a peptide, protein, sense RNA, or ribozyme.

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(Previously Presented): The recombinant DNA molecule of claim 1, wherein

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- 43. (Previously Presented): The recombinant DNA molecule of claim 1, wherein the first regulatory sequence confers endothelium-specific expression *in vivo* of the heterologous DNA sequence.
  - 44. (Previously Presented): An isolated cell transformed with the vector of claim 17.